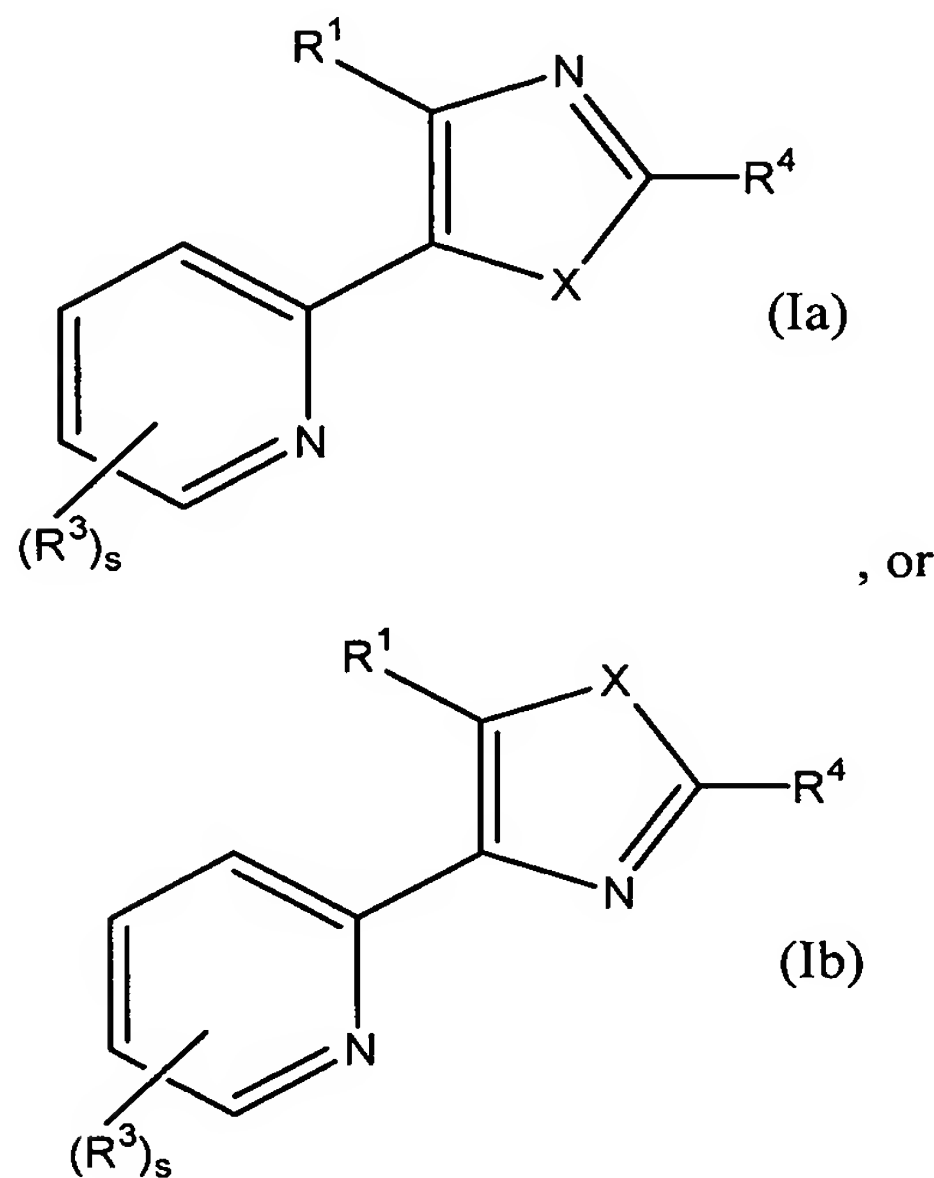


AMENDMENTS TO THE CLAIMS

1. (ORIGINAL) A compound of formula (Ia) or (Ib):



or a pharmaceutically acceptable salt, prodrug, hydrate, tautomer or solvate thereof, wherein:

X is O or S;

R<sup>1</sup> is a saturated, unsaturated, or aromatic C<sub>3</sub>-C<sub>20</sub> mono-, bi- or polycyclic ring optionally containing at least one heteroatom selected from the group consisting of N, O and S, wherein R<sup>1</sup> can optionally be further independently substituted with at least one moiety independently selected from the group consisting of: carbonyl, halo, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkyl, perhalo(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>2</sub>-C<sub>6</sub>)alkenyl, (C<sub>2</sub>-C<sub>6</sub>)alkynyl, hydroxy, oxo, mercapto, (C<sub>1</sub>-C<sub>6</sub>)alkylthio, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>5</sub>-C<sub>10</sub>)aryl or (C<sub>5</sub>-C<sub>10</sub>)heteroaryl, (C<sub>5</sub>-C<sub>10</sub>)aryloxy or (C<sub>5</sub>-C<sub>10</sub>)heteroaryloxy, (C<sub>5</sub>-C<sub>10</sub>)ar(C<sub>1</sub>-C<sub>6</sub>)alkyl or (C<sub>5</sub>-C<sub>10</sub>)heteroar(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>5</sub>-C<sub>10</sub>)ar(C<sub>1</sub>-C<sub>6</sub>)alkoxy or (C<sub>5</sub>-C<sub>10</sub>)heteroar(C<sub>1</sub>-C<sub>6</sub>)alkoxy, HO-(C=O)-, ester, amido, ether, amino, amino(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkylamino(C<sub>1</sub>-C<sub>6</sub>)alkyl, di(C<sub>1</sub>-C<sub>6</sub>)alkylamino(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>5</sub>-C<sub>10</sub>)heterocyclyl(C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl- and di(C<sub>1</sub>-C<sub>6</sub>)alkylamino, cyano, nitro, carbamoyl, (C<sub>1</sub>-C<sub>6</sub>)alkylcarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxycarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl, di(C<sub>1</sub>-C<sub>6</sub>)alkylaminocarbonyl, (C<sub>5</sub>-C<sub>10</sub>)arylcabonyl, (C<sub>5</sub>-C<sub>10</sub>)aryloxycarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkylsulfonyl, and (C<sub>5</sub>-C<sub>10</sub>)arylsulfonyl;

each  $R^3$  is independently selected from the group consisting of: hydrogen, halo, halo( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkyl, ( $C_2$ - $C_6$ )alkenyl, ( $C_2$ - $C_6$ )alkynyl, perhalo( $C_1$ - $C_6$ )alkyl, phenyl, ( $C_5$ - $C_{10}$ )heteroaryl, ( $C_5$ - $C_{10}$ )heterocyclic, ( $C_3$ - $C_{10}$ )cycloalkyl, hydroxy, ( $C_1$ - $C_6$ )alkoxy, perhalo( $C_1$ - $C_6$ )alkoxy, phenoxy, ( $C_5$ - $C_{10}$ )heteroaryl-O-, ( $C_5$ - $C_{10}$ )heterocyclic-O-, ( $C_3$ - $C_{10}$ )cycloalkyl-O-, ( $C_1$ - $C_6$ )alkyl-S-, ( $C_1$ - $C_6$ )alkyl-SO<sub>2</sub>-, ( $C_1$ - $C_6$ )alkyl-NH-SO<sub>2</sub>-, O<sub>2</sub>N-, NC-, amino, Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, ( $C_1$ - $C_6$ )alkyl HN-, ( $C_1$ - $C_6$ )alkylamino, [( $C_1$ - $C_6$ )alkyl]<sub>2</sub>-amino, ( $C_1$ - $C_6$ )alkyl-SO<sub>2</sub>-NH-, amino(C=O)-, aminoO<sub>2</sub>S-, ( $C_1$ - $C_6$ )alkyl-(C=O)-NH-, ( $C_1$ - $C_6$ )alkyl-(C=O)-[(( $C_1$ - $C_6$ )alkyl)-N]-, phenyl-(C=O)-NH-, phenyl-(C=O)-[(( $C_1$ - $C_6$ )alkyl)-N]-, ( $C_1$ - $C_6$ )alkyl-(C=O)-, phenyl-(C=O)-, ( $C_5$ - $C_{10}$ )heteroaryl-(C=O)-, ( $C_5$ - $C_{10}$ )heterocyclic-(C=O)-, ( $C_3$ - $C_{10}$ )cycloalkyl-(C=O)-, HO-(C=O)-, ( $C_1$ - $C_6$ )alkyl-O-(C=O)-, H<sub>2</sub>N(C=O)-, ( $C_1$ - $C_6$ )alkyl-NH-(C=O)-, [( $C_1$ - $C_6$ )alkyl]<sub>2</sub>-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-[( $C_1$ - $C_6$ )alkyl)-N]-(C=O)-, ( $C_5$ - $C_{10}$ )heteroaryl-NH-(C=O)-, ( $C_5$ - $C_{10}$ )heterocyclic-NH-(C=O)-, ( $C_3$ - $C_{10}$ )cycloalkyl-NH-(C=O)- and ( $C_1$ - $C_6$ )alkyl-(C=O)-O-,

where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl, alkoxy, phenoxy, amino of  $R^3$  is optionally substituted by at least one substituent independently selected from ( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkoxy, halo( $C_1$ - $C_6$ )alkyl, halo, H<sub>2</sub>N-, Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, and ( $C_1$ - $C_6$ )alkylHN-;

s is an integer from one to five;

$R^4$  is independently selected from the group consisting of: hydrogen, halo, halo( $C_1$ - $C_6$ )alkyl, ( $C_1$ - $C_6$ )alkyl, ( $C_2$ - $C_6$ )alkenyl, ( $C_2$ - $C_6$ )alkynyl, perhalo( $C_1$ - $C_6$ )alkyl, phenyl, ( $C_5$ - $C_{10}$ )heteroaryl, ( $C_5$ - $C_{10}$ )heterocyclic, ( $C_3$ - $C_{10}$ )cycloalkyl, hydroxy, ( $C_1$ - $C_6$ )alkoxy, perhalo( $C_1$ - $C_6$ )alkoxy, phenoxy, ( $C_5$ - $C_{10}$ )heteroaryl-O-, ( $C_5$ - $C_{10}$ )heterocyclic-O-, ( $C_3$ - $C_{10}$ )cycloalkyl-O-, ( $C_1$ - $C_6$ )alkyl-S-, ( $C_1$ - $C_6$ )alkyl-SO<sub>2</sub>-, ( $C_1$ - $C_6$ )alkyl-NH-SO<sub>2</sub>-, O<sub>2</sub>N-, NC-, amino, Ph(CH<sub>2</sub>)<sub>1-6</sub>HN-, ( $C_1$ - $C_6$ )alkylHN-, ( $C_1$ - $C_6$ )alkylamino, [( $C_1$ - $C_6$ )alkyl]<sub>2</sub>-amino, ( $C_1$ - $C_6$ )alkyl-SO<sub>2</sub>-NH-, amino(C=O)-, aminoO<sub>2</sub>S-, ( $C_1$ - $C_6$ )alkyl-(C=O)-NH-, ( $C_1$ - $C_6$ )alkyl-(C=O)-(( $C_1$ - $C_6$ )alkyl)-N-, phenyl-(C=O)-NH-, phenyl-(C=O)-(( $C_1$ - $C_6$ )alkyl)-N]-, ( $C_1$ - $C_6$ )alkyl-(C=O)-, phenyl-(C=O)-, ( $C_5$ - $C_{10}$ )heteroaryl-(C=O)-, ( $C_5$ - $C_{10}$ )heterocyclic-(C=O)-, ( $C_3$ - $C_{10}$ )cycloalkyl-(C=O)-, HO-(C=O)-, ( $C_1$ - $C_6$ )alkyl-O-(C=O)-, H<sub>2</sub>N(C=O)-, ( $C_1$ - $C_6$ )alkyl-NH-(C=O)-, (( $C_1$ - $C_6$ )alkyl)<sub>2</sub>-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-(( $C_1$ - $C_6$ )alkyl)-N]-(C=O)-, ( $C_5$ - $C_{10}$ )heteroaryl-NH-

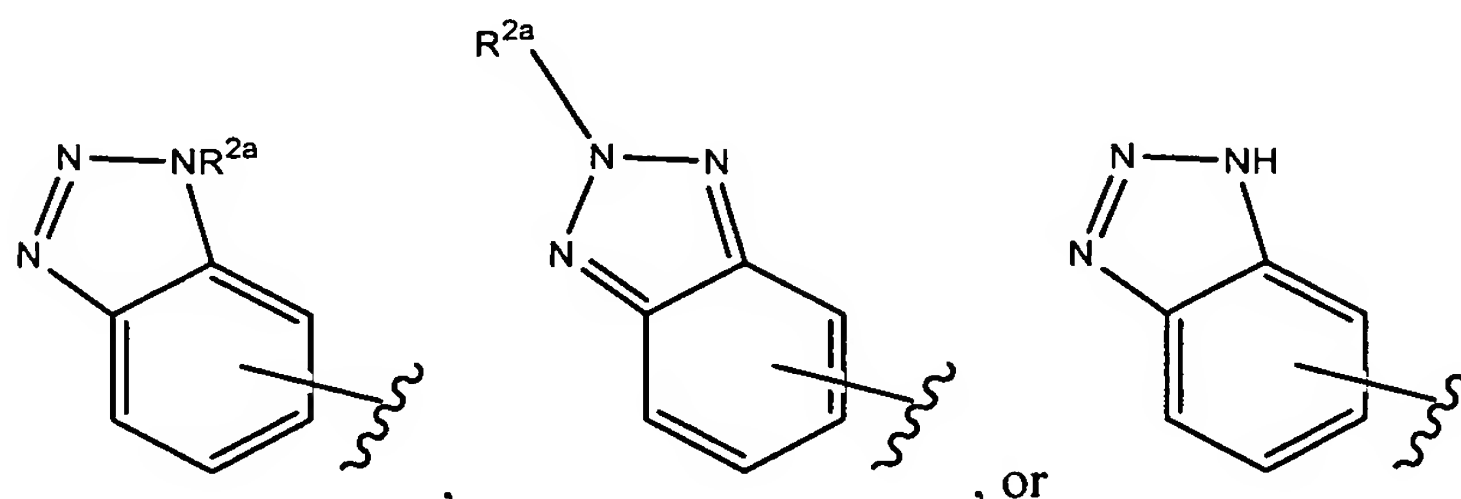
(C=O)-, (C<sub>5</sub>-C<sub>10</sub>)heterocyclic-NH-(C=O)-, (C<sub>3</sub>-C<sub>10</sub>)cycloalkyl-NH-(C=O)- and (C<sub>1</sub>-C<sub>6</sub>)alkyl-(C=O)-O-,

where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl, alkoxy, phenoxy, amino of R<sup>4</sup> is optionally substituted by at least one substituent independently selected from the group consisting of (C<sub>1</sub>-C<sub>6</sub>)alkyl, (C<sub>1</sub>-C<sub>6</sub>)alkoxy, halo(C<sub>1</sub>-C<sub>6</sub>)alkyl, halo, H<sub>2</sub>N-, Ph (CH<sub>2</sub>)<sub>1-6</sub>HN-, (C<sub>1</sub>-C<sub>6</sub>)alkylHN-, (C<sub>5</sub>-C<sub>10</sub>)heteroaryl and (C<sub>5</sub>-C<sub>10</sub>)heterocyclyl; with the proviso that when R<sup>4</sup> is a substituted phenyl moiety, then (a) R<sup>1</sup> is not naphthyl, phenyl or anthracenyl and (b) if R<sup>1</sup> is a phenyl fused with an aromatic or non-aromatic cyclic ring of 5-7 members wherein said cyclic ring optionally contains up to three heteroatoms independently selected from N, O and S, then the fused cyclic ring of said R<sup>1</sup> moiety is substituted;

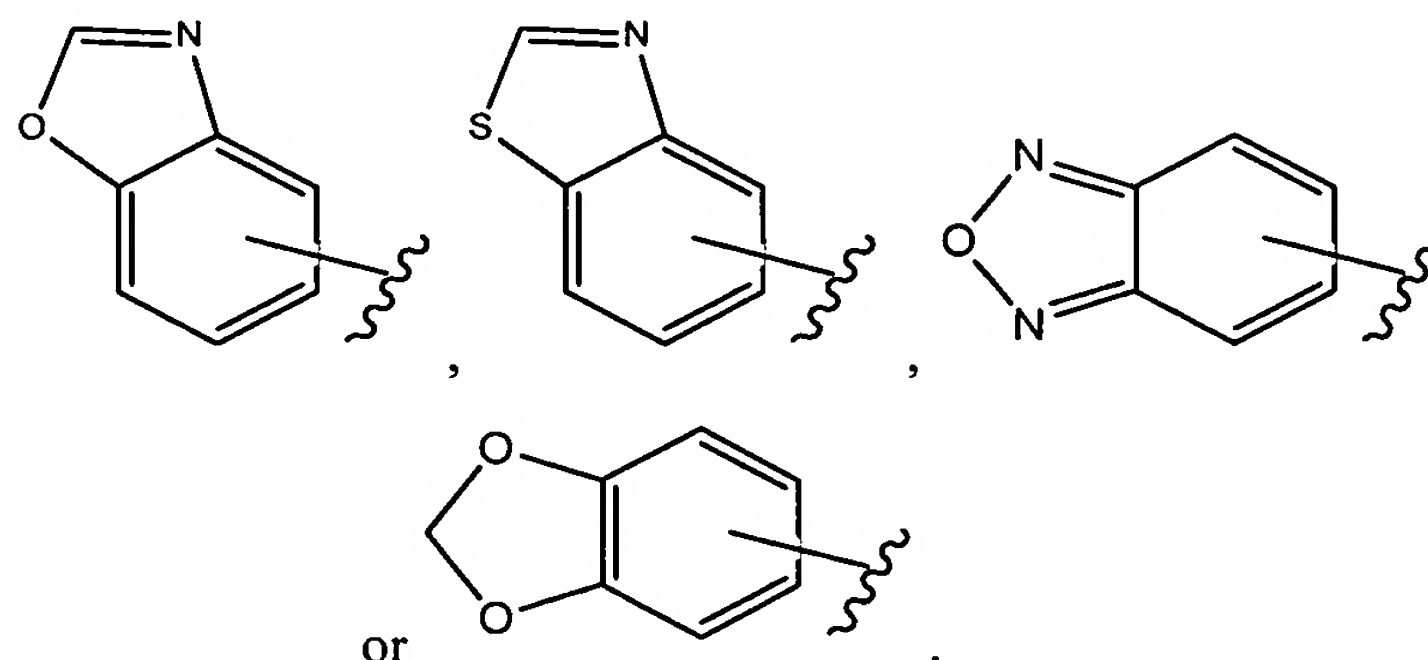
with the proviso that when R<sup>4</sup> is NH<sub>2</sub> and X is S, then R<sup>1</sup> is not an amino-substituted pyridyl or pyrimidinyl moiety; and

with the proviso that when in formula (Ia) R<sup>4</sup> is CH<sub>3</sub> and X is S, R<sup>1</sup> is not a 3, 4-dimethoxy substituted phenyl moiety.

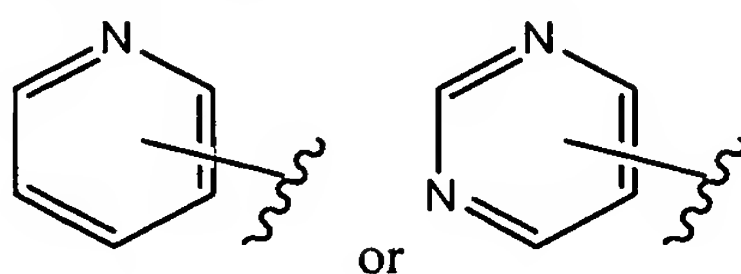
2. (ORIGINAL) A compound of claim 1, wherein R<sup>1</sup> is



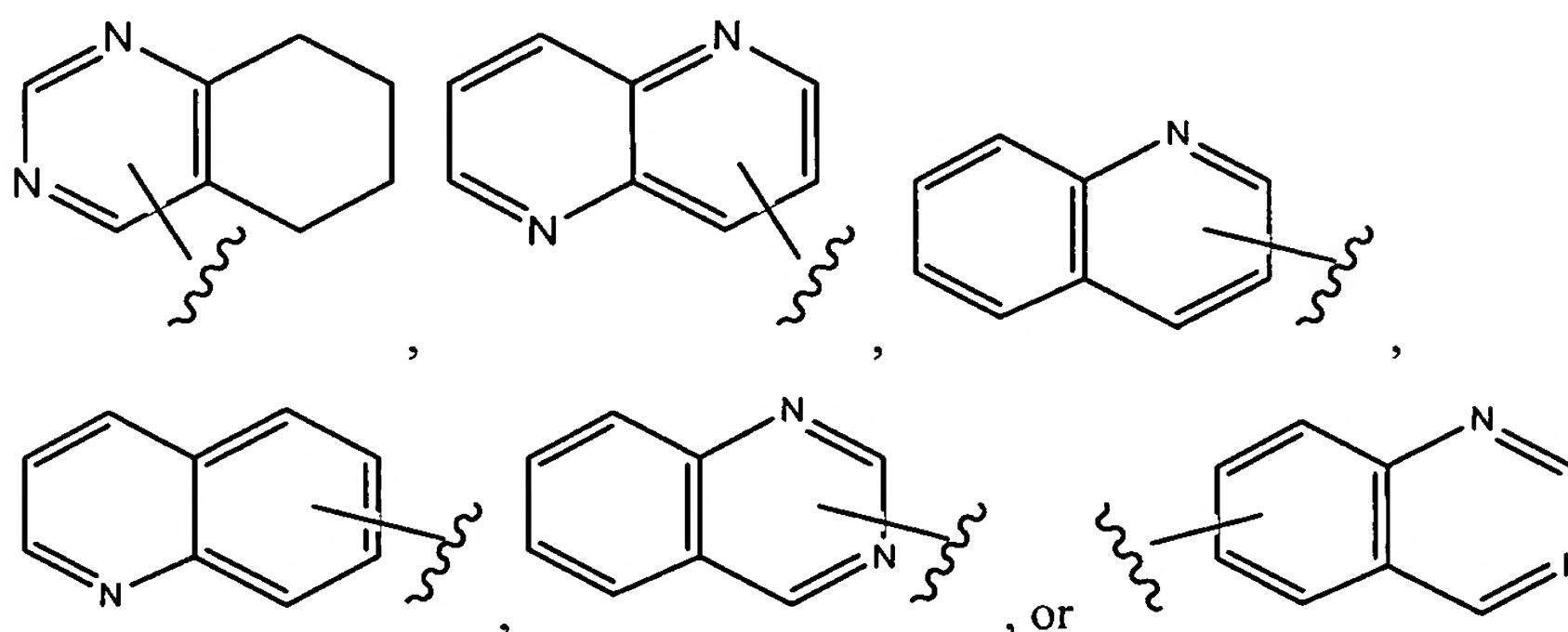
3. (ORIGINAL) A compound of claim 1, wherein R<sup>1</sup> is



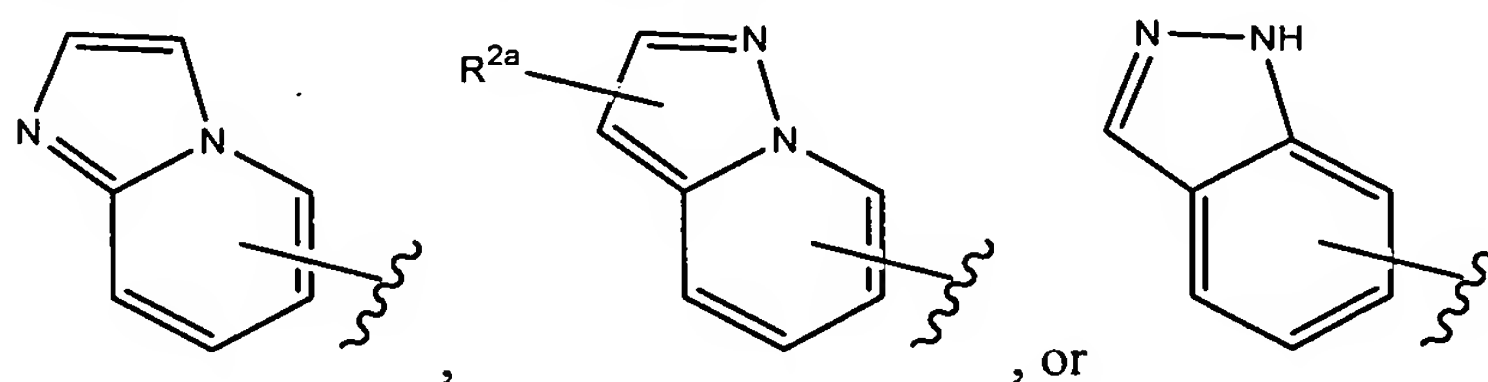
4. (ORIGINAL) A compound of claim 1, wherein  $R^1$  is



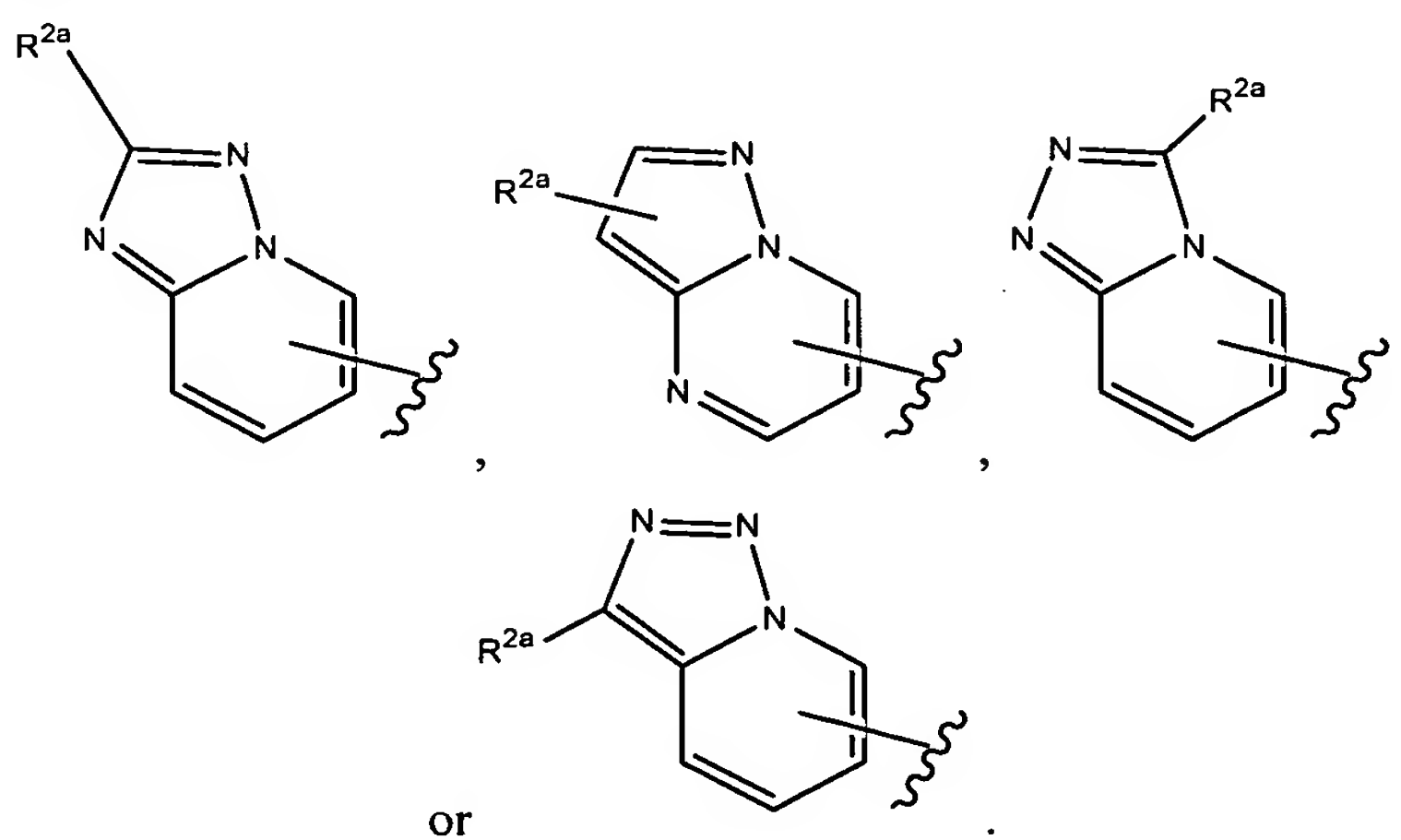
5. (ORIGINAL) A compound of claim 1, wherein  $R^1$  is



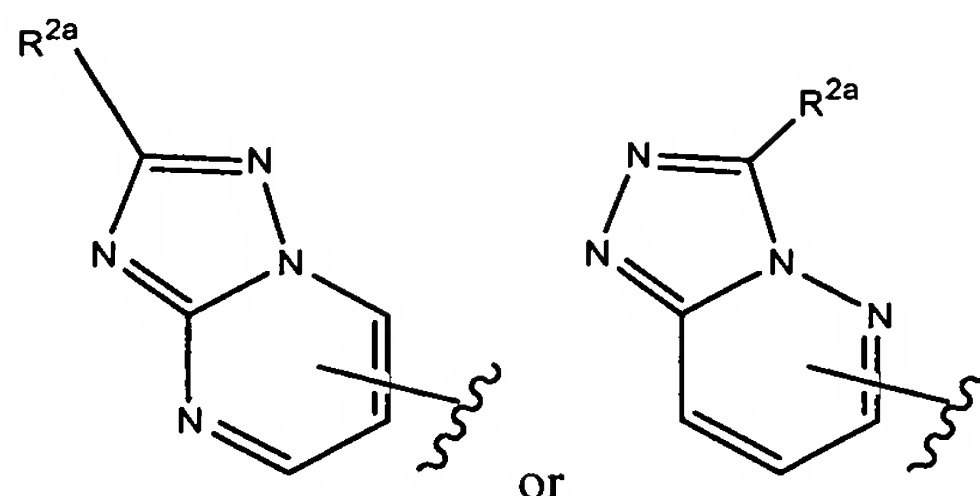
6. (ORIGINAL) A compound of claim 1, wherein  $R^1$  is



7. (ORIGINAL) A compound of claim 1, wherein  $R^1$  is



8. (ORIGINAL) A compound of claim 1, wherein  $R^1$  is



9. (ORIGINAL) A compound of claim 1, wherein X is O; s is one to two;  $R^3$  is hydrogen or  $(C_1-C_6)$ alkyl; and  $R^4$  is H,  $(C_1-C_6)$ alkyl, or amino.

10. (ORIGINAL) A compound of claim 1, wherein X is S; s is one to two;  $R^3$  is hydrogen or  $(C_1-C_6)$ alkyl; and  $R^4$  is H,  $(C_1-C_6)$ alkyl, or amino.

11. (ORIGINAL) A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier.

12. (CURRENTLY AMENDED) A method of ~~preventing or~~ treating a TGF-related disease state in an animal or human comprising the step of administering a therapeutically effective amount of a compound of claim 1 to the animal or human suffering from the TGF-related disease state selected from the group consisting of cancer, glomerulonephritis, diabetic nephropathy, hepatic fibrosis, pulmonary fibrosis, intimal hyperplasia and restenosis, scleroderma, and dermal scarring.

13. (CANCELED)